

In re: Shirley *et al.*

Appl. No.: 09/187,661

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A1
concluded in a concentration of at least about 250 mg/ml, wherein said variant is a polypeptide that has IGF-I activity and differs from the amino acid sequence for said human IGF-I by up to 10 amino acid residues.

Please cancel claim 2 without prejudice or disclaimer.

Please amend claim 3, as follows:

A2
2. (amended) The [IGF-I] composition of claim [2]1, wherein said [syrup comprises] human IGF-I or variant thereof is present in a concentration of about 250 mg/ml to about 500 mg/ml.

Please amend claim 4, as follows:

A3
3. (amended) The [IGF-I] composition of claim [2]1, wherein said [syrup comprises] human IGF-I or variant thereof is present in a concentration of about 350 mg/ml, and wherein said [syrup] composition has a density of about 1.07 g/ml and a viscosity of about 15,700 cps.

Please cancel claims 8 and 12 without prejudice or disclaimer.

Please amend claim 13, as follows:

A4
4. (amended) A kit for reconstituting a pharmaceutical composition comprising biologically active human IGF-I or biologically active variant thereof, said kit comprising [a carrier being compartmentalized to receive in close confinement therein one or more container means, wherein one of said container means contains a highly concentrated form of biologically active IGF-I or variant thereof, wherein said IGF-I or variant thereof is present in a concentration of at least about 250 mg/ml,] the composition of claim 1 and [another of said container means contains] a pharmaceutically acceptable buffered solution having a pH less than or equal to about pH 5.0.

Please amend claim 16, as follows:

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16. (amended) A [composition prepared according to the method of claim 15, wherein said composition is a] pharmaceutical composition comprising the composition of claim 1.

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17. (amended) A [composition prepared according to the method of claim 15, wherein said composition is a]cryogenically produced PLGA microsphere comprising the composition of claim 1.

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18. (amended) The microsphere of claim 17, wherein said microsphere comprises a lyophilized form of said [syrup]composition.

Please add the following new claims:

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19. The composition of claim 1, wherein said variant differs from the amino acid sequence for said human IGF-I by up to 5 amino acid residues.

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20. The composition of claim 1, wherein said variant differs from the amino acid sequence for said human IGF-I by up to 2 amino acid residues.

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21. The composition of claim 1, wherein said variant differs from the amino acid sequence for said human IGF-I by 1 amino acid residue.

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22. The composition of claim 1, wherein said human IGF-I is recombinant human IGF-I.

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23. The composition of claim 13, wherein said recombinant human IGF-I is present at a concentration of about 250 mg/ml to about 500 mg/ml.

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~~38.~~

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The composition of claim ~~34~~, wherein said recombinant human IGF-I is present in a concentration of about 350 mg/ml, and wherein said composition has a density of about 1.07 g/ml and a viscosity of about 15,700 cps.

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~~34.~~

A low salt-containing composition comprising biologically active human IGF-I in a concentration of at least about 250 mg/ml.

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The composition of claim ~~34~~, wherein said human IGF-I is recombinant human IGF-I.

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~~38.~~

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The composition of claim ~~34~~, wherein said human IGF-I is present in a concentration of about 250 mg/ml to about 500 mg/ml.

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The composition of claim ~~34~~, wherein said human IGF-I is present in a concentration of about 350 mg/ml, and wherein said composition has a density of about 1.07 g/ml and a viscosity of about 15,700 cps.

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~~38.~~

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A pharmaceutical composition comprising the composition of claim ~~34~~.

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~~38.~~

A cryogenically produced PLGA microsphere comprising the composition of claim ~~34~~.

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A pharmaceutical composition comprising the composition of claim ~~37~~.

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~~41.~~

A cryogenically produced PLGA microsphere comprising the composition of claim ~~37~~.

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